PAIRITI COULDINALION LIREALI From the INTERNATIONAL SEARCHING AUTHORITY STANLEY J. DAVIS JEREMY M. BEN-DAVID & CO. LTD. P.O. BOX 45087 WRITTEN OPINION OF THE HAR HOTZVIM HI-TECH PARK INTERNATIONAL SEARCHING AUTHORITY JERUSALEM, ISRAEL 91450 (PCT Rule 43bis.1) 1 JAN ZUU Date of mailing (day/month/year) Applicant's or agent's file reference FOR FURTHER ACTION See paragraph 2 below NEU202-13.2 International application No. International filing date (day/month/year) Priority date (day/month/year) PCT/IL04/00567 24 June 2004 (24.06.2004) 26 June 2003 (26.06.2003) International Patent Classification (IPC) or both national classification and IPC IPC(7): C07C 225/00; C07C 67/02 and US Cl.: 564/251; 560/251; 514/551,646 Applicant **NEURIM PHARMACEUTICALS (1991) LTD.** 1. This opinion contains indications relating to the following items: Box No. I Basis of the opinion Box No. II Priority Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability Box No. IV Lack of unity of invention Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement Box No. VI Certain documents cited Box No. VII Certain defects in the international application Box No. VIII Certain observations on the international application

### 2. FURTHER ACTION

If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis (b) that written opinions of this International Searching Authority will not be so considered.

If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later.

For further options, see Form PCT/ISA/220.

3. For further details, see notes to Form PCT/ISA/220.

Name and mailing address of the ISA/ US
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Form PCT/ISA/237 (cover sheet) (January 2004)

## WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.
PCT/IL04/00567

Box No. I Basis of this opinion	
<ol> <li>With regard to the language, this opinion has been established on the basis of the international application in the language it was filed, unless otherwise indicated under this item.</li> </ol>	ge in which
This opinion has been established on the basis of a translation from the original language into the following language which is the language of a translation furnished for the purposes of international search (under Rules 12.3 and 23).	
2. With regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessic claimed invention, this opinion has been established on the basis of:	ary to the
a. type of material	
a sequence listing	
table(s) related to the sequence listing	
b. format of material	
in written format	
in computer readable form	
c. time of filing/furnishing	
contained in international application as filed.	
filed together with the international application in computer readable form.	
furnished subsequently to this Authority for the purposes of search.	
3. In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto filed or furnished, the required statements that the information in the subsequent or additional copies is identical the application as filed or does not go beyond the application as filed, as appropriate, were furnished.	
4. Additional comments:	
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# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.
PCT/IL04/00567

Box N	o. III Non-establishment of opinion with	regard to novelty, inventive step and industrial applicability							
	questions whether the claimed invention appears strially applicable have not been examined in rec	to be novel, to involve an inventive step (to be non-obvious), or to be spect of:							
	the entire international application								
	claims Nos. 11								
beca	use:								
$\boxtimes$	the said international application, or the said claim Nos. 11 relate to the following subject matter which does not require an international preliminary examination (specify):								
	•	The recitation of a use, without setting forth any steps involved in the process, i.e., results in a claim which is not a proper process claim.							
the description, claims or drawings (indicate particular elements below) or said claims Nos are so meaningful opinion could be formed (specify):									
•									
	the claims, or said claims Nos are so in be formed.	nadequately supported by the description that no meaningful opinion could							
	no international search report has been established for said claims Nos.								
	the nucleotide and/or amino acid sequence listing does not comply with the standard provided for in Annex C of the Administrative Instructions in that:								
	the written form	has not been furnished							
		does not comply with the standard							
	the computer readable form	has not been furnished							
		does not comply with the standard							
		o acid sequence listing, if in computer readable form only, do not comply Annex C-bis of the Administrative Instructions.							

Form PCT/ISA/237 (Box No. III) (January 2004)

### WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No. PCT/IL04/00567

Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement 1. Statement Novelty (N) Claims 2-8 YES Claims 1,9-10 NO YES Inventive step (IS) Claims 2-8 Claims 1,9-10 NO Industrial applicability (IA) YES Claims 1-10 Claims NONE NO

#### 2. Citations and explanations:

Claims 1 and 9-10 lack novelty under PCT Article 33(2) and lack an inventive step under PCT Article 33(3) as being anticipated by GB 1334884 and Sm Chem. Abstract 1972:474963.

Both prior art references teach compounds that are fully embraced by claims 1, 9 and 10.

Claims 2-8 meet the criteria set out in PCT Article 33(2)-(3), because the prior art does not teach or fairly suggest the claimed compounds. The compounds of these claims were not taught or suggested by the prior art.

Claims 1-10 meet the criteria set out in PCT Article 33(4), and thus have industrial applicability because the subject matter claimed can be made or used in industry.

Form PCT/ISA/237 (Box No. V) (January 2004)

### WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/IL04/00567

Box No. V	/III	Certain	observations	on the	international	application
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The following observations on the clarity of the claims, description, and drawings or-on the questions whether the claims are fully

supported by the description, are made: Claims 2-4 and 7-8 are objected to under PCT Rule 66.2(a)(v) as lacking clarity under PCT Article 6 because the claims are indefinite for the following reason(s): These claims do not fall within the broad definition of variable X as defined in claim 1. X in claims 2-4 and 7-8 is defined as a C1-C6 alkylene, -C=O, -C=S or a single bond. Claims 2-4 and 7-8 have compounds or definitions outside the scope of the definition of X as indicated in claim 1. Thus the claims are broader than the base claim and render indefinite the metes and bounds of the claims. inglet .